

## SEQUENCE LISTING

<110> Genetics Institute

Karen, Lyons

Vicki, Rosen

Aaron, Daluiski

Thomas, Engstrand

Matthew, Bahamonde

Gamer, Laura Agius, Eric

<120> Method and Composition for Modulating Bone Growth

<130> 22058-554 UTIL

<140> 10/005,228

<141> 2001-12-03

<150> US 60/250,535

<151> 2000-12-01

<160> 9

<170> PatentIn version 3.2

<210> 1

<211> 2952

<212> DNA

<213> Homo sapiens

<400> 1

gaagcgaata gcgttttcag agatattggg cggctcaagg gtcttactct gtcgccagt 60

ctgtaatgca gtgctgtgac catagcccac tgcagcctcc acctcccagg ctcaagcagt 120

cettecece tegeceteat gaatagetgg gactacagee tggageattg gtaagegtea 180

cactgccaaa gtgagagctg ctggagaact cataatccca ggaacgcctc ttctactctc 240

cgagtacccc agtgaccaga gtgagagaag ctctgaacga gggcacgcgg cttgaaggac 300

tgtgggcaga tgtgaccaag agcctgcatt aagttgtaca atggtagatg gagtgatgat 360

tcttcctgtg cttatcatga ttgctctccc ctcccctagt atggaagatg agaagcccaa 420

ggtcaacccc aaactctaca tgtgtgtgtg tgaaggtctc tcctgcggta atgaggacca 480

ctgtgaaggc cagcagtgct tttcctcact gagcatcaac gatggcttcc acgtctacca 540 gaaaggctgc ttccaggttt atgagcaggg aaagatgacc tgtaagaccc cgccgtcccc 600 tggccaaget gtggagtget gccaagggga ctggtgtaac aggaacatca cggcccaget 660 gcccactaaa ggaaaatcct tccctggaac acagaatttc cacttggagg ttggcctcat 720 tattctctct gtagtgttcg cagtatgtct tttagcctgc ctgctgggag ttgctctccg 780 aaaatttaaa aggcgcaacc aagaacgcct caatccccga gacgtggagt atggcactat 840 cgaagggctc atcaccacca atgttggaga cagcacttta gcagatttat tggatcattc 900 gtgtacatca ggaagtggct ctggtcttcc ttttctggta caaagaacag tqqctcqcca 960 gattacactg ttggagtgtg tcgggaaagg caggtatggt gaggtgtgga ggggcagctq 1020 gcaaggggaa aatgttgccg tgaagatctt ctcctcccgt gatgagaagt catggttcag 1080 ggaaacggaa ttgtacaaca ctgtgatgct gaggcatgaa aatatcttag gtttcattgc 1140 ttcagacatg acatcaagac actccagtac ccagctgtgg ttaattacac attatcatga 1200 aatgggatcg ttgtacgact atcttcagct tactactctg gatacagtta gctgccttcg aatagtgctg tccatagcta gtggtcttgc acatttgcac atagagatat ttgggaccca 1320 agggaaacca gccattgccc atcgagattt aaagagcaaa aatattctgg ttaagaagaa 1380 tggacagtgt tgcatagcag atttgggcct ggcagtcatg cattcccaga gcaccaatca 1440 gcttgatgtg gggaacaatc cccgtgtggg caccaagcgc tacatggccc ccgaagttct 1500 agatgaaacc atccaggtgg attgtttcga ttcttataaa agggtcgata tttgggcctt 1560

tggacttgtt ttgtgggaag tggccaggcg gatggtgagc aatggtatag tggaggatta 1620 caagccaccg ttctacgatg tggttcccaa tgacccaagt tttgaagata tgaggaaggt 1680 agtctgtgtg gatcaacaaa ggccaaacat acccaacaga tggttctcag acccgacatt 1740 aacctctctg gccaagctaa tgaaagaatg ctggtatcaa aatccatccg caagactcac 1800 agcactgcgt atcaaaaaga ctttgaccaa aattgataat tccctcgaca aattgaaaac 1860 tgactgttga cattttcata gtgtcaagaa ggaagatttg acgttgttgt cattgtccag 1920 ctgggaccta atgctggcct gactggttgt cagaatggaa tccatctgtc tccctccca 1980 aatggctgct ttgacaaggc agacgtcgta cccagccatg tgttggggag acatcaaaac 2040 caccctaacc tegetegatg actgtgaact gggcatttca egaactgtte acactgcaga 2100 gactaatgtt ggacagacac tgttgcaaag gtagggactg gaggaacaca gagaaatcct 2160 aaaagagatc tgggcattaa gtcagtggct ttgcatagct ttcacaagtc tcctagacac 2220 tccccacggg aaactcaagg aggtggtgaa tttttaatca gcaatattgc ctgtgcttct 2280 cttctttatt gcactaggaa ttctttgcat tccttacttg cactgttact cttaatttta aagacccaac ttgccaaaat gttggctgcg tactccactg gtctgtcttt ggataatagg 2400 aattcaattt ggcaaaacaa aatgtaatgt cagactttgc tgcattttac acatgtgctg 2460 atgtttacaa tgatgccgaa cattaggaat tgtttataca caactttgca aattatttat tacttgtgca cttagtagtt tttacaaaac tgctttgtgc atatgttaaa gcttattttt 2580 atgtggtctt atgattttat tacagaaatg tttttaacac tatactctaa aatggacatt 2640

ttcttttatt atcagttaaa atcacatttt aagtgcttca catttgtatg tgtgtagact 2700

gtaacttttt ttcagttcat atgcagaacg tatttagcca ttacccacgt gacaccaccg 2760

aatatattat cgatttagaa gcaaagattt cagtagaatt ttagtcctga acgctacggg 2820

gaaaatgcat tttcttcaga attatccatt acgtgcattt aaactctgcc agaaaaaaat 2880

tgttttcaag tc 2952

<210> 2

<211> 509

<212> PRT

<213> Homo sapiens

<400> 2

Met Val Asp Gly Val Met Ile Leu Pro Val Leu Ile Met Ile Ala Leu 1 5 10 15

Pro Ser Pro Ser Met Glu Asp Glu Lys Pro Lys Val Asn Pro Lys Leu 20 25 30

Tyr Met Cys Val Cys Glu Gly Leu Ser Cys Gly Asn Glu Asp His Cys 35 40 45

Glu Gly Gln Gln Cys Phe Ser Ser Leu Ser Ile Asn Asp Gly Phe His
50 55 60

Val Tyr Gln Lys Gly Cys Phe Gln Val Tyr Glu Gln Gly Lys Met Thr 65 70 75 80

Cys Lys Thr Pro Pro Ser Pro Gly Gln Ala Val Glu Cys Cys Gln Gly
85 90 95

Asp Trp Cys Asn Arg Asn Ile Thr Ala Gln Leu Pro Thr Lys Gly Lys 100 105 110

Ser Phe Pro Gly Thr Gln Asn Phe His Leu Glu Val Gly Leu Ile Ile Leu Ser Val Val Phe Ala Val Cys Leu Leu Ala Cys Leu Leu Gly Val Ala Leu Arg Lys Phe Lys Arg Arg Asn Gln Glu Arg Leu Asn Pro Arg Asp Val Glu Tyr Gly Thr Ile Glu Gly Leu Ile Thr Thr Asn Val Gly Asp Ser Thr Leu Ala Asp Leu Leu Asp His Ser Cys Thr Ser Gly Ser Gly Ser Gly Leu Pro Phe Leu Val Gln Arg Thr Val Ala Arg Gln Ile Thr Leu Leu Glu Cys Val Gly Lys Gly Arg Tyr Gly Glu Val Trp Arg Gly Ser Trp Gln Gly Glu Asn Val Ala Val Lys Ile Phe Ser Ser Arg Asp Glu Lys Ser Trp Phe Arg Glu Thr Glu Leu Tyr Asn Thr Val Met Leu Arg His Glu Asn Ile Leu Gly Phe Ile Ala Ser Asp Met Thr Ser Arg His Ser Ser Thr Gln Leu Trp Leu Ile Thr His Tyr His Glu Met Gly Ser Leu Tyr Asp Tyr Leu Gln Leu Thr Thr Leu Asp Thr Val Ser Cys Leu Arg Ile Val Leu Ser Ile Ala Ser Gly Leu Ala His Leu His Ile Glu Ile Phe Gly Thr Gln Gly Lys Pro Ala Ile Ala His Arg Asp

325	330	335
~ _ ~	<b>550</b>	222

Leu Lys Ser Lys Asn Ile Leu Val Lys Lys Asn Gly Gln Cys Cys Ile 340 345 350

Ala Asp Leu Gly Leu Ala Val Met His Ser Gln Ser Thr Asn Gln Leu 355 360 365

Asp Val Gly Asn Asn Pro Arg Val Gly Thr Lys Arg Tyr Met Ala Pro 370 375 380

Glu Val Leu Asp Glu Thr Ile Gln Val Asp Cys Phe Asp Ser Tyr Lys 385 390 395 400

Arg Val Asp Ile Trp Ala Phe Gly Leu Val Leu Trp Glu Val Ala Arg 405 410 415

Arg Met Val Ser Asn Gly Ile Val Glu Asp Tyr Lys Pro Pro Phe Tyr 420 425 430

Asp Val Val Pro Asn Asp Pro Ser Phe Glu Asp Met Arg Lys Val Val 435 440 445

Cys Val Asp Gln Gln Arg Pro Asn Ile Pro Asn Arg Trp Phe Ser Asp 450 455 460

Pro Thr Leu Thr Ser Leu Ala Lys Leu Met Lys Glu Cys Trp Tyr Gln 465 470 475 480

Asn Pro Ser Ala Arg Leu Thr Ala Leu Arg Ile Lys Lys Thr Leu Thr 485 490 495

Lys Ile Asp Asn Ser Leu Asp Lys Leu Lys Thr Asp Cys 500 505

<210> 3

<211> 1969

<212> DNA

<213> Homo sapiens

<400> 3

aggaaacggt ttattaggag ggagtggtgg agctgggcca ggcaggaaga cgctggaata 60 agaaacattt ttgctccagc ccccatccca gtcccgggag gctgccgcgc cagctgcgcc 120 gagcgagccc ctccccggct ccagcccggt ccggggccgc gccggacccc agcccgccgt 180 ccagcgctgg cggtgcaact gcggccgcgc ggtggagggg aggtggcccc ggtccgccga 240 aggetagege ceegeeacee geagageggg ceeagaggga ceatgaeett gggeteecee 300 aggaaaggee ttetgatget getgatggee ttggtgaeee agggagaeee tgtgaageeg 360 tctcggggcc cgctggtgac ctgcacgtgt gagagcccac attgcaaggg gcctacctgc cggggggcct ggtgcacagt agtgctggtg cgggaggagg ggaggcaccc ccaggaacat 480 cggggctgcg ggaacttgca cagggagctc tgcagggggc gccccaccga gttcqtcaac 540 cactactget gegacageea cetetgeaae cacaaegtgt ceetggtget ggaggeeaee 600 caacctcctt cggagcagcc gggaacagat ggccagctgg ccctgatcct gggccccgtg 660 ctggccttgc tggccctggt ggccctgggt gtcctgggcc tgtggcatgt ccgacggagg 720 caggagaagc agcgtggcct gcacagcgag ctgggagagt ccaqtctcat cctqaaaqca 780 tetgageagg gegaeacgat gttgggggae eteetggaea gtgaetgeae eacagggagt ggeteaggge teecetteet ggtgeagagg acagtggeae ggeaggttge ettggtggag 900 tgtgtgggaa aaggccgcta tggcgaagtg tggcggggct tgtggcacgg tgagagtgtg 960 gccgtcaaga tcttctcctc gagggatgaa cagtcctggt tccgggagac tgagatctat 1020 aacacagtat tgctcagaca cgacaacatc ctaggcttca tcgcctcaga catgacctcc 1080

cgcaactcga gcacgcagct gtggctcatc acgcactacc acgagcacgg ctccctctac 1140

gactttctgc agagacagac gctggagccc catctggctc tgaggctagc tgtgtccgcg 1200

gcatgcggcc tggcgcacct gcacgtggag atcttcggta cacagggcaa accagccatt 1260

cccaccgcga cttcaagagc cgcaatgtgc tggtcaagag caacctgcag tgttgcatcg 1320

ccgacctggg cctggctgtg atgcactcac agggcagcga ttacctggac atcggcaaca 1380

accegagagt gggcaccaag eggtacatgg caccegaggt getggaegag cagateegea 1440

cggactgctt tgagtcctac aagtggactg acatctgggc ctttggcctg gtgctgtggg 1500

agattgcccg ccggaccatc gtgaatggca tcgtggagga ctatagacca cccttctatg 1560

atgtggtgcc caatgacccc agctttgagg acatgaagaa ggtggtgtgt gtggatcagc 1620

agacccccac catccctaac cggctggctg cagacccggt cctctcaggc ctagctcaga 1680

tgatgcggga gtgctggtac ccaaacccct ctgcccgact caccgcgctg cggatcaaga 1740

agacactaca aaaaattagc aacagtccag agaagcctaa agtgattcaa tagcccagga 1800

gcacctgatt cctttctgcc tgcagggggc tgggggggtg gggggcagtg gatggtgccc 1860

tatctgggta gaggtagtgt gagtgtggtg tgtgctgggg atgggcagct gcgcctgcct 1920

gctcggcccc cagcccaccc agccaaaaat acagctgggc tgaaacctg 1969

<210> 4

<211> 503

<212> PRT

<213> Homo sapiens

<400> 4

Met Thr Leu Gly Ser Pro Arg Lys Gly Leu Leu Met Leu Leu Met Ala

1

Leu Val Thr Gln Gly Asp Pro Val Lys Pro Ser Arg Gly Pro Leu Val 20 25 30

Thr Cys Thr Cys Glu Ser Pro His Cys Lys Gly Pro Thr Cys Arg Gly 35 40 45

Ala Trp Cys Thr Val Val Leu Val Arg Glu Glu Gly Arg His Pro Gln 50 55 60

Glu His Arg Gly Cys Gly Asn Leu His Arg Glu Leu Cys Arg Gly Arg 65 70 75 80

Pro Thr Glu Phe Val Asn His Tyr Cys Cys Asp Ser His Leu Cys Asn 85 90 95

His Asn Val Ser Leu Val Leu Glu Ala Thr Gln Pro Pro Ser Glu Gln 100 105 110

Pro Gly Thr Asp Gly Gln Leu Ala Leu Ile Leu Gly Pro Val Leu Ala 115 120 125

Leu Leu Ala Leu Val Ala Leu Gly Val Leu Gly Leu Trp His Val Arg
130 135 140

Arg Arg Gln Glu Lys Gln Arg Gly Leu His Ser Glu Leu Gly Glu Ser 145 150 155 160

Ser Leu Ile Leu Lys Ala Ser Glu Gln Gly Asp Thr Met Leu Gly Asp 165 170 175

Leu Leu Asp Ser Asp Cys Thr Thr Gly Ser Gly Ser Gly Leu Pro Phe 180 185 190

Leu Val Gln Arg Thr Val Ala Arg Gln Val Ala Leu Val Glu Cys Val 195 200 205

Gly Lys Gly Arg Tyr Gly Glu Val Trp Arg Gly Leu Trp His Gly Glu 210 215 220

Ser 225	Val	Ala	Val	Lys	Ile 230	Phe	Ser	Ser	Arg	Asp 235	Glu	Gln	Ser	Trp	Phe 240
Arg	Glu	Thr	Glu	Ile 245	Tyr	Asn	Thr	Val	Leu 250	Leu	Arg	His	Asp	Asn 255	Ile
Leu	Gly	Phe	Ile 260	Ala	Ser	Asp	Met	Thr 265	Ser	Arg	Asn	Ser	Ser 270	Thr	Gln
Leu	Trp	Leu 275	Ile	Thr	His	Tyr	His 280	Glu	His	Gly	Ser	Leu 285	Tyr	Asp	Phe
Leu	Gln 290	Arg	Gln	Thr	Leu	Glu 295	Pro	His	Leu	Ala	Leu 300	Arg	Leu	Ala	Val
Ser 305	Ala	Ala	Cys	Gly	Leu 310	Ala	His	Leu	His	Val 315	Glu	Ile	Phe	Gly	Thr 320
Gln	Gly	Lys	Pro	Ala 325	Ile	Ala	His	Arg	Asp 330	Phe	Lys	Ser	Arg	Asn 335	Val
Leu	Val	Lys	Ser 340	Asn	Leu	Gln	Cys	Cys 345	Ile	Ala	Asp	Leu	Gly 350	Leu	Ala
Val	Met	His 355	Ser	Gln	Gly	Ser	Asp 360	Tyr	Leu	Asp	Ile	Gly 365	Asn	Asn	Pro
Arg	Val 370	Gly	Thr	Lys	Arg	Tyr 375	Met	Ala	Pro	Glu	Val 380	Leu	Asp	Glu	Gln
Ile 385	Arg	Thr	Asp	Cys	Phe 390	Glu	Ser	Tyr	Lys	Trp 395	Thr	Asp	Ile	Trp	Ala 400
Phe	Gly	Leu	Val	Leu 405	Trp	Glu	Ile	Ala	Arg 410	Arg	Thr	Ile	Val	Asn 415	Gly
Ile	Val	Glu	Asp 420	Tyr	Arg	Pro	Pro	Phe 425	Tyr	Asp	Val	Val	Pro 430	Asn	Asp
Pro	Ser	Phe	Glu	Asp	Met	Lys	Lys	Val	Val	Cys	Val	Asp	Gln	Gln	Thr

435 440 445

Pro Thr Ile Pro Asn Arg Leu Ala Ala Asp Pro Val Leu Ser Gly Leu 450 455 460

Ala Gln Met Met Arg Glu Cys Trp Tyr Pro Asn Pro Ser Ala Arg Leu 465 470 475 480

Thr Ala Leu Arg Ile Lys Lys Thr Leu Gln Lys Ile Ser Asn Ser Pro 485 490 495

Glu Lys Pro Lys Val Ile Gln 500

<210> 5

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: signal peptide

<400> 5

Met Pro Leu Leu Leu Leu Leu Leu Leu Pro Ser Pro Leu His Pro 1 5 10 15

<210> 6

<211> 1774

<212> DNA

<213> Homo sapiens

<400> 6

agatettgaa aacaceeggg ceacacaege egegaeetae agetetttet eagegttgga 60

gtggagacgg cgcccgcagc gccctgcgcg ggtgaggtcc gcgcagctgc tggggaagag 120

cccacctgtc aggctgcgct gggtcagcgc agcaagtggg gctggccgct atctcgctgc 180

acceggeege gteeeggget eegtgegeee tegeeeeage tggtttggag tteaaccete 240

ggctccgccg ccggctcctt gcgccttcgg agtgtcccgc agcgacgccg ggagccgacg 300

cgccgcgcgg gtacctagcc atggctgggg cgagcaggct gctctttctg tggctgggct 360 gcttctgcgt gagcctggcg cagggagaga gaccgaagcc acctttcccg gagctccgca 420 aagetgtgee aggtgaeege aeggeaggtg gtggeeegga eteegagetg eageegeaag 480 acaaggtete tgaacacatg etgeggetet atgacaggta cagcacggte caggeggee ggacaccggg ctccctggag ggaggctcgc agccctggcg ccctcggctc ctgcgcgaag 600 gcaacacggt tcgcagcttt cgggcggcag cagcagaaac tcttgaaaga aaaggactgt 660 atatcttcaa tctgacatcg ctaaccaagt ctgaaaacat tttgtctgcc acactgtatt tctgtattgg agagctagga aacatcagcc tgagttgtcc agtgtctgga ggatgctccc 780 atcatgctca gaggaaacac attcagattg atctttctgc atggaccctc aaattcagca 840 gaaaccaaag tcaactcctt ggccatctgt cagtggatat ggccaaatct catcgagata 900 ttatgtcctg gctgtctaaa gatatcactc aactcttgag gaaggccaaa gaaaatgaag 960 agttcctcat aggatttaac attacgtcca agggacgcca gctgccaaag aggaggttac 1020 cttttccaga gccttatatc ttggtatatg ccaatgatgc cgccatttct gagccagaaa 1080 gtgtggtatc aagcttacag ggacaccgga attttcccac tggaactgtt cccaaatggg atagccacat cagagctgcc ctttccattg agcggaggaa gaagcgctct actggggtct 1200 tgctgcctct gcagaacaac gagcttcctg gggcagaata ccagtataaa aaggatgagg 1260 tgtgggagga gagaaagcct tacaagaccc ttcaggctca ggcccctgaa aagagtaaga 1320 ataaaaagaa acagagaaag gggcctcatc ggaagagcca gacgctccaa tttgatgagc 1380

agaccctgaa aaaggcaagg agaaagcagt ggattgaacc tcggaattgc gccaggagat 1440

acctcaaggt agactttgca gatattggct ggagtgaatg gattatctcc cccaagtcct 1500

ttgatgccta ttattgctct ggagcatgcc agttccccat gccaaagtct ttgaagccat 1560

caaatcatgc taccatccag agtatagtga gagctgtggg ggtcgttcct gggattcctg 1620

agccttgctg tgtaccagaa aagatgtcct cactcagtat tttattcttt gatgaaaata 1680

agaatgtagt gcttaaagta taccctaaca tgacagtaga gtcttgcgct tgcagataac 1740

ctggcaaaga actcatttga atgcttaatt caat 1774

<210> 7

<211> 472

<212> PRT

<213> Homo sapiens

<400> 7

Met Ala Gly Ala Ser Arg Leu Leu Phe Leu Trp Leu Gly Cys Phe Cys 1 5 10 15

Val Ser Leu Ala Gln Gly Glu Arg Pro Lys Pro Pro Phe Pro Glu Leu 20 25 30

Arg Lys Ala Val Pro Gly Asp Arg Thr Ala Gly Gly Gly Pro Asp Ser 35 40 45

Glu Leu Gln Pro Gln Asp Lys Val Ser Glu His Met Leu Arg Leu Tyr 50 55 60

Asp Arg Tyr Ser Thr Val Gln Ala Ala Arg Thr Pro Gly Ser Leu Glu 65 70 75 80

Gly Gly Ser Gln Pro Trp Arg Pro Arg Leu Leu Arg Glu Gly Asn Thr 85 90 95

Val Arg Ser Phe Arg Ala Ala Ala Glu Thr Leu Glu Arg Lys Gly

Leu '	Tyr	Ile 115	Phe	Asn	Leu	Thr	Ser 120	Leu	Thr	Lys	Ser	Glu 125	Asn	Ile	Leu
Ser i	Ala 130	Thr	Leu	Tyr	Phe	Cys 135	Ile	Gly	Glu	Leu	Gly 140	Asn	Ile	Ser	Leu
Ser (	Cys	Pro	Val	Ser	Gly 150	Gly	Cys	Ser	His	His 155	Ala	Gln	Arg	Lys	His 160
Ile (	Gln	Ile	Asp	Leu 165	Ser	Ala	Trp	Thr	Leu 170	Lys	Phe	Ser	Arg	Asn 175	Gln
Ser (	Gln	Leu	Leu 180	Gly	His	Leu	Ser	Val 185	Asp	Met	Ala	Lys	Ser 190	His	Arg
Asp 1	Ile	Met 195	Ser	Trp	Leu	Ser	Lys 200	Asp	Ile	Thr	Gln	Leu 205	Leu	Arg	Lys
Ala I	Lys 210	Glu	Asn	Glu	Glu	Phe 215	Leu	Ile	Gly	Phe	Asn 220	Ile	Thr	Ser	Lys
Gly 1 225	Arg	Gln	Leu	Pro	Lys 230	Arg	Arg	Leu	Pro	Phe 235	Pro	Glu	Pro	Tyr	Ile 240
Leu V	Val	Tyr	Ala	Asn 245	Asp	Ala	Ala	Ile	Ser 250	Glu	Pro	Glu	Ser	Val 255	Val
Ser S	Ser	Leu	Gln 260	Gly	His	Arg	Asn	Phe 265	Pro	Thr	Gly	Thr	Val 270	Pro	Lys
Trp A	Asp	Ser 275	His	Ile	Arg	Ala	Ala 280	Leu	Ser	Ile	Glu	Arg 285	Arg	Lys	Lys
Arg S	Ser 290	Thr	Gly	Val	Leu	Leu 295	Pro	Leu	Gln	Asn	Asn 300	Glu	Leu	Pro	Gly
Ala (	Glu	Tyr	Gln	Tyr	Lys 310	Lys	Asp	Glu	Val	Trp 315	Glu	Glu	Arg	Lys	Pro 320

Tyr Lys Thr Leu Gln Ala Gln Ala Pro Glu Lys Ser Lys Asn Lys Lys 325 330 335 Lys Gln Arg Lys Gly Pro His Arg Lys Ser Gln Thr Leu Gln Phe Asp 340 345 350 Glu Gln Thr Leu Lys Lys Ala Arg Arg Lys Gln Trp Ile Glu Pro Arg Asn Cys Ala Arg Arg Tyr Leu Lys Val Asp Phe Ala Asp Ile Gly Trp 370 375 380 Ser Glu Trp Ile Ile Ser Pro Lys Ser Phe Asp Ala Tyr Tyr Cys Ser 385 390 395 400 Gly Ala Cys Gln Phe Pro Met Pro Lys Ser Leu Lys Pro Ser Asn His 405 410 Ala Thr Ile Gln Ser Ile Val Arg Ala Val Gly Val Val Pro Gly Ile 420 425 Pro Glu Pro Cys Cys Val Pro Glu Lys Met Ser Ser Leu Ser Ile Leu 440 435 445 Phe Phe Asp Glu Asn Lys Asn Val Val Leu Lys Val Tyr Pro Asn Met 450 455 460 Thr Val Glu Ser Cys Ala Cys Arg 465 <210> 8 <211> 17 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: RT-PCR Primer <400> ggacagacgc tgctatt

17

```
<210> 9
<211> 17
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: RT-PCR Primer
<400> 9
tgttctacga ctcactc
17
```

. . . . .